



**UNITED STATES PATENT AND TRADEMARK OFFICE**

**Applicant(s):** Mark A. Dombroski et al.

**Examiner:** Evelyn Mei Huang

**Serial No:** 10/649,227

**Art Unit:** 1625

**Filed:** August 27, 2003

**Docket:** 17474 (PC25308A)

**For:** ALKYL-[4-(DIFLUOROPHENYL)-  
OXAZOL-5-YL]-TRIAZOLO-  
PYRIDINES

**Confirmation No.: 5400**

Commissioner for Patents  
United States Patent and Trademark Office  
P.O. Box 1450  
Alexandria, Virginia 22313-1450

**DECLARATION OF DR. KIM F. MCCLURE**  
**UNDER 37 C.F.R. §1.132**

Sir:

I, Kim F. McClure, hereby declare as follows:

1. I am an applicant of U.S. Application Serial No.10/649,227, filed August 27, 2003, which claims the benefit of U.S. Serial No. 60/407,088, filed August 30, 2002;
2. I hold a Doctorate Degree in the field of Chemistry from Yale University which I obtained in 1993;
3. I have been employed at Pfizer, Inc. since 1995, and my current position is Senior Principal Scientist;
4. A true and correct copy of my Curriculum Vitae is enclosed herein as Exhibit A;

5. I have reviewed the above-identified application (hereinafter referred to as '227 application), and U.S. Patent No. 6,696,464 B2 (hereinafter referred to as the '464 patent) and I am familiar with the subject matter therein;

6. It is my scientific opinion that the two closest structural compounds between the '227 application and the '464 patent are 6-[4-(4-Fluoro-phenyl)-oxazol-5-yl]-3-isopropyl-[1,2,4]triazolo[4,3-a]pyridine (Example 12) in the '464 patent and 3-tert-Butyl-6-[4-(2,4-difluoro-phenyl)-oxazol-5-yl]-[1,2,4]triazolo[4,3-a]pyridine (Example 4) in the '227 application;

7. 6-[4-(4-Fluoro-phenyl)-oxazol-5-yl]-3-isopropyl-[1,2,4]triazolo[4,3-a]pyridine has a human whole blood TNF alpha IC<sub>50</sub> value of 685 nM;

8. 3-tert-Butyl-6-[4-(2,4-difluoro-phenyl)-oxazol-5-yl]-[1,2,4]triazolo[4,3-a]pyridine has a human whole blood TNF alpha IC<sub>50</sub> value of 76 nM;

9. It is my scientific opinion that the human whole blood TNF alpha IC<sub>50</sub> value of 76 nM for 3-tert-Butyl-6-[4-(2,4-difluoro-phenyl)-oxazol-5-yl]-[1,2,4]triazolo[4,3-a]pyridine shows that this compound is a surprisingly and unexpectedly better TNF alpha inhibitor than 6-[4-(4-Fluoro-phenyl)-oxazol-5-yl]-3-isopropyl-[1,2,4]triazolo[4,3-a]pyridine, which has a human whole blood TNF alpha IC<sub>50</sub> of 685 nM;

10. I declare that all statements made herein of my own knowledge are true and that all statements are believed to be true; and that those statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code, and that such willful false statements may jeopardize the validity of the application or any patent issuing thereon.

By: Kim F McClure  
Dr. Kim F. McClure

Dated: October 1, 2004

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## Vita for Kim F. McClure

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### • EDUCATION

B.S. Chemistry, May 1988. University of California, Berkeley

Ph. D. Chemistry, December 1993. Yale University

### • EXPERIENCE

Pfizer Inc., 1995-present. Senior Principal Scientist

Post-doctoral Research Fellow (NIH), 1993-present, M.I.T. Advisor: Daniel S. Kemp  
— Synthesis and study of  $\alpha$ -helix templates and their peptide conjugates

Graduate Research Fellow, 1988-1993, Yale University. Advisor: Samuel J. Danishefsky  
— Synthesis of FR-900482 congeners. Solid-phase carbohydrate synthesis using glycals.

Undergraduate research, 1987-88, U. C. Berkeley. Advisor: Clayton H. Heathcock  
— Studies on zinc enolates. Synthesis of Daphnilactone A

Teaching Assistant, 1987-1990, Yale University (4 semesters); U.C.Berkeley (2 semesters)  
— Introductory through graduate organic chemistry courses

### • AWARDS

National Institutes of Health post-doctoral fellowship (1993-present)

Samuel K. Bushnell graduate fellowship (1989-1990)

Department of Education graduate fellowship (1990-1991)RFS

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## PUBLICATIONS

Reiter, Lawrence A.; Robinson, Ralph P.; McClure, Kim F.; Jones, Christopher S.; Reese, Matthew R.; Mitchell, Peter G.; Otterness, Ivan G.; Bliven, Marcia L.; Liras, Jennifer; Cortina, Santo R.; Donahue, Kathleen M.; Eskra, James D.; Griffiths, Richard J.; Lame, Mary E.; Lopez-Anaya, Arturo; Martinelli, Gary J.; McGahee, Shunda M.; Yocum, Sue A.; Lopresti-Morrow, Lori L.; Tobiassen, Lisa M.; Vaughn-Bowser, Marcie L. Pyran-containing sulfonamide hydroxamic acids: potent MMP inhibitors that spare MMP-1. *Bioorganic & Medicinal Chemistry Letters* 2004, 14, 3389-3395.

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McClure, K. F.; Axt, M. Z. *Bioorg. & Med. Chem. Lett.* 1998, 8, 143. Alkylation of Succinates: Synthesis of Ro 32-3555.

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Cammers-Goodwin, A.; Allen, T. J.; Oslick, S. L.; McClure, K. F.; Lee, J. H.; Kemp, D. S. *J. Am. Chem. Soc.* 1996, 118, 3082. Mechanism of Stabilization of Helical Conformations of Polypeptides by Water Containing Trifluoroethanol.

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McClure, K. F.; Danishefsky, S. J. *J. Org. Chem.* 1991 56, 850. Cycloaddition Reactions of Aromatic Nitroso Compounds with Oxygenated Dienes. An Approach to the Synthesis of the FR-900482 Family of Antibiotics.

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Robinson, Ralph Pelton, Jr.; McClure, Kim Francis. Preparation of arylsulfonylaminoalkylhydroxamates as inhibitors of matrix metalloproteinases or tumor necrosis factor production. WO 9833768 A1

• REFERENCES

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